

Claims

- 1 Stable pharmaceutical compositions of 5,10-methylene-(6R)-, -(6S)-or -(6R,S)-tetrahydrofolate, characterised in that the composition comprises 5,10-methylene-(6R)-, -(6S)- or -(6R,S)-tetrahydrofolic acid or a pharmaceutically acceptable salt of 5,10-methylene-(6R)-, -(6S)- or -(6R,S)-tetrahydrofolic acid together with citrate, and has a pH between 7.5 and 10.5, preferably between 8.5 and 9.5.
- 2 Stable pharmaceutical compositions according to claim 1 together with further pharmaceutically acceptable active ingredients and adjuvants.
- 3 A pharmaceutical composition according to claim 2, characterised in that it comprises formaldehyde as an adjuvant.
- 15 4 A pharmaceutical composition according to claim 2, characterised in that it comprises a further folate as a further active ingredient.
- 5 A pharmaceutical composition according to claim 4, characterised in that it comprises tetrahydrofolic acid and salts thereof as a further folate.
- 20 6 A pharmaceutical composition according to claim 1, characterised in that the calcium salt or an acidic salt is used as the pharmaceutically acceptable salt of 5,10-methylene-(6R)-, -(6S)- or -(6R,S)-tetrahydrofolic acid.
- 25 7 A pharmaceutical composition according to claim 2, characterised in that it comprises a cytostatic agent as a further active ingredient .
- 8 A pharmaceutical composition according to claim 2, characterised in that it comprises a fluorinated pyrimidine derivative as a further active ingredient.

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- 9 A pharmaceutical composition according to claim 8, characterised in that it comprises 5-fluoruracil or a 5-fluoruracil prodrug, particularly capecitabine (xeloda) as a fluorinated pyrimidine derivative.
- 5 10 A pharmaceutical composition according to any one of claims 1 to 9, additionally comprising at least one antioxidant or a radical scavenger.
- 11 A pharmaceutical composition according to claim 10, characterised in that it comprises vitamin C or reduced glutathione as an antioxidant or radical scavenger.
- 10 12 A pharmaceutical composition according to any one of claims 1 to 11, characterised in that the composition exists as a lyophilisate, dry powder or dry mixture.
- 13 A pharmaceutical composition according to any one of claims 1 to 11, characterised in 15 that the composition exists as a lyophilisation solution.
- 14 A method of stabilising compositions comprising 5,10-methylene-(6R)-, -(6S)- or -(6R,S)-tetrahydrofolate, characterised in that 5,10-methylene-(6R)-, -(6S)- or -(6R,S)-tetrahydrofolic acid is treated with citrate and is brought to a pH between 7.5 and 10.5, 20 preferably between 8.5 and 9.5.
- 15 Use of compositions comprising a pharmaceutically acceptable salt of 5,10-methylene-(6R)-, -(6S)- or -(6R,S)-tetrahydrofolic acid and citrate at a pH between 7.5 and 10.5, 25 preferably between 8.5 and 9.5, for producing a pharmaceutical preparation suitable for use for corresponding medical indications.